

### AMENDMENTS TO THE CLAIMS

This listing of claims will replace all prior versions, and listings, of claims in the application:

1. (Currently amended) A Method method for ~~the determination of~~ determining the free fraction of a substance comprising
  - (a) ~~incubation of~~ incubating the substance with a suspension of particles, ~~other than erythrocytes~~, having a lipophilic surface, in a ~~substantially~~ protein-free aqueous medium, ~~for the determination of the distribution of the substance between the particles and said substantially protein-free medium~~ wherein the particles and the medium are erythrocyte-free;
  - (b) determining the distribution of the substance between the particles and said protein free medium in step (a);
  - ~~(b)(c)~~ incubation of incubating the substance with a suspension of particles, ~~other than erythrocytes~~, having a lipophilic surface, in a protein-containing aqueous medium, ~~for the determination of the distribution of the substance between the particles and said protein-containing aqueous medium~~ wherein the particles and the medium are erythrocyte-free;
  - (d) determining the distribution of the substance between the particles and said protein-containing aqueous medium in step (c);
  - ~~(e)(c)~~ ~~determination of~~ determining the free fraction of the substance from the distributions determined under steps (a)(b) and ~~(b)(d)~~.
2. (Currently amended) The Method method of claim 1, wherein said ~~suspension of particles is~~ are selected from a group consisting of ~~suspensions comprising~~
  - (a) ~~a suspension of~~ particles having a solid core; <sub>1</sub>
  - ~~(b)~~ ~~a suspension of~~ particles having a solid core comprising a silica bead; <sub>1</sub> and
  - ~~(c)~~ ~~a suspension of~~ Transil<sup>®</sup> particles.
3. (Previously presented) The Method method of claim 1 or 2, wherein the protein-containing aqueous medium is plasma.
4. (Currently amended) The Method method of ~~any of claims 1 to 3~~ claim 1 or 2, wherein the ~~substantially~~ protein-free aqueous medium is a buffer solution.

5. (Currently amended) ~~The Method~~ method of ~~any of claims 1 to 4~~ claim 1 or 2, wherein ~~said incubations of~~ said substance is incubated with said suspensions of particles ~~is~~ on a plate having multiple cavities or on a 96-well-plate.
6. (Currently amended) ~~The Method~~ method of ~~any of claims 1 to 4~~ claim 2, wherein ~~said particles have the solid core~~ is a ferromagnetic solid core.
7. (Currently amended) ~~A Method~~ method for ~~the determination of~~ determining the relative free fraction of a substance in a first species in relation to the free fraction of the same substance in a second species comprising
  - (a) determining the membrane affinity of said substance in plasma ( $MA_{\text{plasma}}$ ) of ~~said substance for~~ said first species,
  - (b) determining the membrane affinity of said substance in plasma ( $MA_{\text{plasma}}$ ) of ~~said substance for~~ said second species,
  - (c) determining the relative free fraction from the results determined under the steps (a) and (b);

provided that the membrane affinity of said substance in plasma ( $MA_{\text{plasma}}$ ) of said first species and said second species is determined in an erythrocyte-free environment.
8. (Withdrawn) A kit for use in any of the methods of claims 1 to 7, comprising a plate having multiple cavities, a buffer solution, plasma, and particles selected from a group of particles comprising
  - (a) particles having a solid core;
  - (b) particles having a solid core which is a silica bead; and
  - (c) Transil<sup>®</sup> particles.
9. (Withdrawn) The kit of claim 8, comprising plasma of two different species.
10. (Withdrawn) A kit of claim 8 or 9, wherein said specific amounts of particles are placed within said cavities of said plate.
11. (New) The method of claim 2, wherein said particles have a solid core coated with lipophilic drugs.